

JS006348069B1

(12) United States Patent

Vacanti et al.

1.995 970 A

4,137,921 A

4,141,087 A

4,144,126 A

4,186,448 A

4,192,827 A

4,205,399 A

4,228,243 A

4,239,664 A

4,243,775 A

(10) Patent No.:

US 6,348,069 B1

(45) Date of Patent:

Feb. 19, 2002

(54)	ENGINEERING OF STRONG, PLIABLE TISSUES						
(75)	Inventors:	Joseph P. Vacanti, Winchester; Christopher K. Breuer, Brighton; Beverly E. Chaignaud; Toshiraru Shin'oka, both of Brookline, all of MA (US)					
(73)	Assignee:	Children's Medical Center Corporation, Boston, MA (US)					
(*)	Notice:	Subject to any disclaimer, the term of this patent is extended or adjusted under 35 U.S.C. 154(b) by 0 days.					
(21)	Appl. No.: 09/185,360						
(22)	Filed:	Nov. 3, 1998					
Related U.S. Application Data							
(62)	Division of application No. 08/445,280, filed on May 19, 1995, now Pat. No. 5,855,610.						
(51)	Int. Cl.7	A61F 2/02					
	U.S. Cl						
	Field of Search						
(56)		References Cited					
U.S. PATENT DOCUMENTS							

4,277,582 A	7/1981	Mueller et al.
4,280,954 A	7/1981	Yannas et al.
4,304,591 A	12/1981	Mueller et al.
4,304,866 A	12/1981	Green et al.
4,328,204 A	5/1982	Wasserman et al.
4,347,847 A	9/1982	Usher
4,348,329 A	9/1982	Chapman
4,352,883 A	10/1982	Lim
4,356,261 A	10/1982	Kuettner
4,391,797 A	7/1983	Folkman et al.
4,416,986 A	11/1983	Markus et al.
4,427,808 A	1/1984	Stol et al.
4,431,428 A	2/1984	Schmer
4,438,198 A	3/1984	Schmer
4,439,152 A	3/1984	Small
4,440,921 A	4/1984	Allcock et al.
4,444,887 A	4/1984	Hoffmann
4,446,229 A	5/1984	Indech
4,446,234 A	5/1984	Russo et al.
4,450,150 A	5/1984	Sidman
4,456,687 A	6/1984	Green
4,458,678 A	7/1984	Yannas et al.
4,485,096 A	11/1984	Bell
4,485,097 A	11/1984	Bell
4,489,056 A	12/1984	Himmelstein et al.
4,494,385 A	1/1985	Kuraoka et al.
4,495,174 A	1/1985	Allcock et al.
4,505,266 A	3/1985	Yannas et al.
4,520,821 A	6/1985	Schmidt et al.
4,528,265 A	7/1985	Becker
4,544,516 A	10/1985	Hughes et al.
4,545,082 A	10/1985	Hood
4,553,272 A	11/1985	Mears
	_	

(List continued on next page.)

FOREIGN PATENT DOCUMENTS

	(List continued on next page.)			
DE	35 18 150	10/1986		
DE.	28 53 614	7/1979		
AU	24245/88 B	2/1989		

OTHER PUBLICATIONS

Allcock & Kwon, "An Ionically Cross-Linkable Polyphosphazene: Poly[bis(carboxylatophenoxy)phosphazene] and Its Hydrogels and Membranes," *Macromolecules* 22:75-79 (1989).

(List continued on next page.)

Primary Examiner—David J Isabella
(74) Attorney, Agent, or Firm—Holland & Knight LLP

(57) ABSTRACT

It has been discovered that improved yields of engineered tissue following implantation, and engineered tissue having enhanced mechanical strength and flexibility or pliability, can be obtained by implantation, preferably subcutaneously, of a fibrous polymeric matrix for a period of time sufficient to obtain ingrowth of fibrous tissue and/or blood vessels, which is the removed for subsequent implantation at the site where the implant is desired. The matrix is optionally seeded prior to the first implantation, after ingrowth of the fibrous tissue, or at the time of reimplantation. The time required for fibrous ingrowth typically ranges from days to weeks. The method is particularly useful in making valves and tubular structures, especially heart valves and blood vessels.

7 Claims, No Drawings

1,773,710	^		3/1933	Dorougii	
2,609,347	Α		9/1952	Wilson	
2,653,917	Α		9/1953	Hammon	
2,659,935	A		11/1953	Hammon	
2,664,366	Α		12/1953	Wilson	
2,676,945	Α		4/1954	Higgins	
2,683,136	Α		7/1954	Higgins	
2,703,316	Α		3/1955	Schneider	
2,758,987	Α		8/1956	Salzberg	
2,846,407	Α		8/1958	Wilson	
2,951,828	Α		9/1960	Zeile et al.	
3,514,791	Α	•	6/1970	Sparks	623/66
3,531,561	Α		9/1970	Trehu	
3,826,241	Α		7/1974	Bucalo	
3,880,991	Α		4/1975	Yolles	
3,883,393	Α		5/1975	Knazek et al.	
3,902,497	Α		9/1975	Casey	
3,935,065	Α		1/1976	Doerig	
3,949,073	Α		4/1976	Daniels et al.	
3,960,150	Α		6/1976	Hussain et al.	
3,974,526	Α		8/1976	Dardik et al.	
3,992,725	Α		11/1976	Homsy	
3,995,444	Α		12/1976	Clark et al.	
4,026,304	Α		5/1977	Levy	
4,060,081	Α		11/1977	Yannas et al.	
4,069,307	Α		1/1978	Higuchi et al.	
4 4 0 5 0 6 4					

2/1979 Okuzumi

3/1979 Burbidge

2/1980 Brekke

10/1980 Iizuka

12/1980 Teag et al.

2/1979 Shalaby et al.

3/1980 Mueller et al.

6/1980 Shalaby et al.

1/1981 Rosensaft et al.

3/1935 Dorough